CLAIMS

1. A compound of the formula (I):

$$\begin{array}{c|c}
R^{5} \\
N \\
R^{2}
\end{array}$$

$$\begin{array}{c|c}
Y & R^{3} \\
\end{array}$$
(I)

5 wherein $\label{eq:condition} \text{X is -CO- or -(CH$_2)$_k$- (wherein k is 1, 2 or 3);}$

Y is

20

25

(1) lower alkyl, or

10 (2) $Z - (CH_2)_n -$

{wherein

Z is

- (1) aryl, or
- (2) R¹-CO-NR⁴-

15 (wherein

R¹ is (1) aryl, heterocyclyl,

aryl-(lower alkyl),

aryl-(lower alkoxy), or

heterocyclyl-(lower alkoxy),

each of which may be substituted

with one or more substituent(s)

selected from the group

consisting of

- (a) lower alkyl,
- (b) halogen and
- (c) hydroxy; or

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(2) lower alkoxy; and
               R4 is hydrogen, or lower alkyl); and
          n is 1, 2, 3, 4, 5 or 6};
   R<sup>2</sup> is (1) lower alkyl, aryl-(lower alkyl) or
5
              (lower alkyl)thio-(lower alkyl),
             each of which may be substituted with one
             or more substituent(s) selected from the
             group consisting of
10
                (a) heterocyclyl,
                (b) carboxy,
                (c) carboxy-(lower alkyl),
                (d) amidated carboxy,
                (e) (lower alkoxy) carbonyl which may be
15
                                   with
                    substituted
                                            cycloalkyl,
                    heterocyclyl or (lower alkanoyl)oxy;
                    and
                (f) cyano; or
          (2) aryl which may be substituted with
20
              lower alkyl, lower alkenyl, aryl,
              lower alkoxy, (lower alkyl)amino,
              (lower alkyl)thio, carboxy,
              (lower alkoxy) carbonyl,
              (lower alkoxy) - (lower alkyl),
25
              (lower alkyl)amino-(lower alkyl), or
              (lower alkyl)thio-(lower alkyl),
              each of which may be further substituted with
              one or more substituent(s) selected from the
              group consisting of
30
                (a) heterocyclyl,
                (b) (lower alkoxy) carbonyl,
                (c) carboxy and
                (d) amidated carboxy;
35 R^3 is (1) -Q-R^7,
```

[wherein Q is -CO- or $-SO_2-$, R⁷ is (a) lower alkyl which may be substituted with one or more substituent (s) selected from the 5 group consisting of cycloalkyl, aryl which may be further substituted with aryl(s), and heterocyclyl, 10 (b) lower alkenyl which may be substituted with one or more substituent(s) selected from the group consisting of aryl heterocyclyl, (c) cycloalkyl, 15 (d) aryl which may be substituted with one or more substituent(s) selected from the group consisting of lower alkyl, ary 1 which may be further substituted with 20 hydroxy(s), lower alkoxy, aryloxy, hydroxy, and halogen, 25 (e) hete rocyclyl which may be substituted with one or more substituent(s) selected from the group consisting of lower alkyl, aryl which may be further substituted with 30 halogen(s), and halogen, (f) aryloxy, or (g) amino which may be substituted with aryl(s) which may be further substituted with one 35 or more substituent(s) selected from the

group consisting of aryl and heterocyclyl];
or

(2) lower alkyl which may be substituted with aryl(s) or heterocyclyl(s), each of which may be further substituted with aryl(s); and

 $\ensuremath{\text{R}}^5$ and $\ensuremath{\text{R}}^6$ are independently hydrogen or lower alkyl; or

10 R^6 and Y may be linked together to form $-(CH_2)_m-(wherein m is 2, 3, 4 or 5);$

5

20

or a pharmaceutically acceptable salt thereof.

15 2. A compound of claim 1 having the formula (Ia):

$$Z \xrightarrow{(CH_2)n} \begin{array}{c} H \\ N \\ R^2 \\ N \\ O \end{array}$$
(Ia)

wherein Z, R^2 , R^7 and n are as defined above.

3. A compound of claim 1 having the formula (Ib):

wherein R^1 , R^2 , R^7 and n are as defined above.

A compound of claim 3,

```
wherein
          R1 is aryl-(lower alkoxy);
          R<sup>2</sup> is lower alky, or
                                                                                                                                    with
                                                               may be
                                                                                                 substituted
                                        which
 5
                         aryl
                         carboxy-(lower alkyl);
          R^7 is heterocyclyl which may be substituted with
                         substituted with lower alkyl; and
          n is 1, 2, 3, 4 or 5.
10
                    A compound selected from:
                                         6-{(2S)-2-[(1-benzofuran-2-y1-carbony1)
           sodium
           amino]-5-[benzyloxycarbonylamino]pentanoylamino}-
           hexanoate,
           (2E)-3-{2-[(2S)-2-[(1H-indol-2-ylcarbonyl)amino]-5
15
           -[benzyloxycarbonylamino]pentanoylamino]phenyl}-
           acrylic acid,
           (2E)-3-\{2-[(2S)-2-[(1-methyl-1H-indol-2-yl-1])]
           carbonyl)amino]-5-[benzyloxycarbonylamino]-
           pentanoylamino]phenyl}acrylic acid,
20
           3-\{2-[(2S)-2-[(1-methyl-1H-indol-2-ylcarbonyl)-
           amino]-5-[benzyloxycarbonylamino]pentanoylamino]-
           phenyl}propanoic acid,
            sodium 3-\{2-[(2S)-2-[(2-quinolinylcarbonyl)amino]-
            5-[benzyloxycarbonylamino]pentanoylamino]phenyl}-
 25
            propanoate,
            6-[((2S)-2-[(1-benzofuran-2-ylcarbonyl)amino]-5-
            {[(benzyloxy)carbonyl]amino}pentanoyl)amino]-2-
            naphthoic acid,
            3 - \{2 - [((2S) - 5 - \{[(benzyloxy) carbonyl] amino\} - 2 - \{[(8 - 2) - 2 - [((2S) - 2) - 2 - [((2S) - 2) - 2 - (((2S) - 2) - 2) - (((2S) - 2) - 
 30
            methylimidazo[1, 2-a]pyridin-2-yl)carbonyl]amino}-
            pentanoy1) amino] phenyl} propanoic acid,
             3-[2-({(2S)-5-{[(benzyloxy)carbonyl]amino}-2-
             [(2-quinolinylmethyl)amino]pentanoyl}amino)-
            phenyl]propanoic acid, and
 35
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3-[2-({(2S)-5-{[(benzyloxy)carbonyl]amino}-2-[(1H-indol-2-ylcarbonyl)amino]pentanoyl}amino)phenyl]-propanoic acid.

5 6. A process for preparing the compound of the formula (Ia-1):

$$\begin{array}{c|c}
R^{5} \\
| \\
N \\
R^{2} \\
NR^{6} \\
| \\
Q \\
R^{7} \\
(I a-1)
\end{array}$$

wherein

Y is

15

20

10 (1) lower alkyl, or

(2) $Z - (CH_2)_n -$,

{wherein

Z is

(1) aryl, or

(2) $R^{1}-CO-NR^{4}-$

(wherein

consisting of

```
(a) lower alkyl,
                            (b) halogen and
                            (c) hydroxy; or
                      (2) lower alkoxy; and
5
                R4 is hydrogen, or lower alkyl); and
           n is 1, 2, 3, 4, 5 or 6};
    Q is -CO- or -SO_2-;
10
    R<sup>2</sup> is (1) lower alkyl, aryl-(lower alkyl) or
              (lower alkyl) thio-(lower alkyl),
              each of which may be substituted with one
              or more substituent(s) selected from the
              group consisting of
15
                (a) heterocyclyl,
                (b) carboxy,
                (c) carboxy-(lower alkyl),
                (d) amidated carboxy,
                (e) (lower alkoxy) carbonyl which may be
20
                    substituted
                                    with
                                             cycloalkyl,
                    heterocyclyl or (lower alkanoyl)oxy;
                    and
                (f) cyano; or
          (2) aryl which may be substituted with
25
              lower alkyl, lower alkenyl, aryl,
              lower alkoxy, (lower alkyl)amino,
              (lower alkyl)thio, carboxy,
              (lower alkoxy) carbonyl,
              (lower alkoxy) - (lower alkyl),
30
              (lower alkyl) amino-(lower alkyl), or
              (lower alkyl) thio-(lower alkyl),
              each of which may be further substituted with
              one or more substituent(s) selected from the
              group consisting of
35
                (a) heterocyclyl,
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(b) (lower alkoxy)carbonyl,

```
(c) carboxy and
                (d) amidated carboxy;
    R<sup>5</sup> and R<sup>6</sup> are independently hydrogen or lower alkyl;
5
    or
    R^6 and Y may be linked together to form - (CH_2)_m- (wherein
      m is 2, 3, 4 or 5); and
10
    R<sup>7</sup> is (a) lower alkyl which may be substituted with
             one or more substituent(s) selected from the
             group consisting of
                 cycloalkyl,
                  aryl which may be further substituted with
15
                 aryl(s), and
                 heterocyclyl,
          (b) lower alkenyl which may be substituted with
             one or more substituent(s) selected from the
20
             group consisting of aryl and heterocyclyl,
          (c) cycloalkyl,
          (d) aryl which may be substituted with one or
             more substituent(s) selected from the group
             consisting of
25
                  lower alkyl,
                  aryl which may be further substituted with
                  hydroxy(s),
                  lower alkoxy,
                  aryloxy,
30
                  hydroxy, and
                  halogen,
           (e) heterocyclyl which may be substituted with
             one or more substituent(s) selected from the
             group consisting of
35
                  lower alkyl,
```

aryl which may be further substituted with halogen(s), and halogen,

- (f) aryloxy, or
- 5 (g) amino which may be substituted with aryl(s) which may be substituted with one or more substituent(s) selected from the group consisting of aryl and heterocyclyl];
- 10 or a pharmaceutically acceptable salt thereof,

comprising, reacting a compound (IIa):

(wherein Y and R^6 are each as defined above), or its reactive derivative at the carboxy group or the salt thereof, with a compound (IIIa):

$$HN \stackrel{R^2}{\underset{R^5}{=}}$$

(wherein R^2 and R^5 are each as defined above), or its reactive derivative at the amino group or the salt thereof to give a compound (IVa):

$$\begin{array}{c|c}
R^5 \\
| \\
N \\
R^2 \\
N \\
N \\
R^6 \\
H \\
(IVa)
\end{array}$$

(wherein Y, R^2 , R^5 and R^6 are each as defined above), or its salt; and reacting the compound (IVa):

$$\begin{array}{cccc}
R^5 \\
| \\
N \\
R^2 \\
N \\
R^6 \\
H
\end{array}$$
(IVa)

5

(wherein Y, R^2 , R^5 and R^6 are each as defined above), or its salt, with a compound (V):

$$R^7$$
 OH (V)

(wherein Q and R^7 are each as defined above), or its 10 reactive derivative at the carboxy group (in case of Q is -CO-)/the sulfo group (in case of Q is -SO₂-), or the salt thereof.

7. A process for preparing the compound of the formula 15 (Ib-1):

wherein

15

25

X is -CO-, or -(CH₂)_k- (wherein k is 1, 2 or 3);

5 Q is $-CO- or -SO_2-;$

each of which may be substituted with one or more substituent(s) selected from the group consisting of

- (a) lower alkyl,
- (b) halogen and
- (c) hydroxy; or
- (2) lower alkoxy; and
- R² is (1) lower alkyl, aryl-(lower alkyl) or (lower alkyl) thio-(lower alkyl),

each of which may be substituted with one or more substituent(s) selected from the group consisting of

- (a) heterocyclyl,
- (b) carboxy,
- (c) carboxy-(lower alkyl),
 - (d) amidated carboxy,

```
(e) (lower alkoxy) carbonyl which may be
                                              cycloalkyl,
                    substituted
                                      with
                    heterocyclyl or (lower alkanoyl)oxy;
                     and
                 (f) cyano; or
5
          (2) aryl which may be substituted with
               lower alkyl, lower alkenyl, aryl,
               lower alkoxy, (lower alkyl)amino,
               (lower alkyl)thio, carboxy,
               (lower alkoxy) carbonyl,
10
               (lower alkoxy) - (lower alkyl),
               (lower alkyl)amino-(lower alkyl), or
               (lower alkyl) thio-(lower alkyl),
               each of which may be further substituted with
               one or more substituent(s) selected from the
15
               group consisting of
                 (a) heterocyclyl,
                 (b) (lower alkoxy) carbonyl,
                 (c) carboxy and
                 (d) amidated carboxy;
20
     R<sup>5</sup> and R<sup>6</sup> are independently hydrogen or lower alkyl;
     οr
     R^6 and Y may be linked together to form - (CH_2)_m- (wherein
25
       m is 2, 3, 4 or 5);
     R<sup>7</sup> is (a) lower alkyl which may be substituted with
              one or more substituent(s) selected from the
              group consisting of
30
                  cycloalkyl,
                  aryl which may be further substituted with
                  aryl(s), and
                  heterocycly1,
           (b) lower alkenyl which may be substituted with
35
```

one or more substituent(s) selected from the group consisting of aryl and heterocyclyl, (c) cycloalkyl, (d) aryl which may be substituted with one or 5 more substituent(s) selected from the group consisting of lower alkyl, aryl which may be further substituted with hydroxy(s), 10 lower alkoxy, aryloxy, hydroxy, and halogen, (e) heterocyclyl which may be substituted with 15 one or more substituent(s) selected from the group consisting of lower alkyl, aryl which may be further substituted with halogen(s), and 20 halogen, (f) aryloxy, or (q) amino which may be substituted with aryl(s) which may be substituted with one or more substituent(s) selected from the group 25 consisting of aryl and heterocyclyl]; and n is 1, 2, 3, 4, 5 or 6; or a pharmaceutically acceptable salt thereof, 30 comprising, reacting a compound (IIb):

$$R^{5}$$
 N
 R^{2}
 N
 R^{6}
 R^{6}

(IIb)

(wherein X, R^2 , R^5 , R^6 and n are each as defined above), or its reactive derivative at the amino group or the salt thereof, with a compound (IIIb):

5

(wherein R^1 is as defined above), or its reactive derivative at the carboxy group or the salt thereof to give a compound (IVb):

10

(wherein X, R^1 , R^2 , R^5 , R^6 , n and are as defined above), or its salt; and reacting the compound (IVb):

(wherein X, R^1 , R^2 , R^5 , R^6 and n are as defined above), or its salt, with a compound (V):

$$\mathbb{R}^7$$
 OH \mathbb{Q}

- 5 (wherein Q and R^7 are as defined above), or its reactive derivative at the carboxy group (in case of Q is -CO-)/the sulfo group (in case of Q is -SO₂-), or the salt thereof.
- 10 8. A process for preparing the compound of the formula (Ia-2):

O
$$\mathbb{R}^{2^{\prime}}$$
 OH $\mathbb{R}^{2^{\prime}}$ OH $\mathbb{R}^{2^{\prime}}$ OH $\mathbb{R}^{2^{\prime}}$ (I a-2)

wherein Y is

```
(1) lower alkyl, or
        (2) Z-(CH_2)_n-,
          {wherein
            Zis
 5
             (1) aryl, or
             (2) R<sup>1</sup>-CO-NR<sup>4</sup>-
                (wherein
                 R<sup>1</sup> is (1) aryl, heterocyclyl,
                           aryl-(lower alkyl),
10
                           aryl-(lower alkoxy), or
                           heterocyclyl-(lower alkoxy),
                           each of which may be substituted
                           with one or more substituent(s)
                           selected from
                                               the
                                                       group
15
                           consisting of
                              (a) lower alkyl,
                             (b) halogen and
                             (c) hydroxy; or
                       (2) lower alkoxy; and
20
               . R4 is hydrogen, or lower alkyl); and
           n is 1, 2, 3, 4, 5 or 6};
    Q is -CO- or -SO_2-;
25
    R<sup>2</sup> is (1) lower alkyl, (lower alkyl) thio-(lower alkyl)
              or aryl-(lower alkyl); or
           (2) aryl which may be substituted with
              lower alkyl, lower alkenyl, aryl,
              lower alkoxy, (lower alkyl)amino,
30
              (lower alkyl)thio,
              (lower alkoxy) - (lower alkyl),
              (lower alkyl) amino-(lower alkyl), or
              [(lower alkyl)thio]-(lower alkyl);
35
   R<sup>6</sup> is hydrogen or lower alkyl; or
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 \mbox{R}^{6} and Y may be linked together to form $-\left(\mbox{CH}_{2}\right)_{\,m}-$ (m is

2, 3, 4 or 5); ${ t R}^7$ is (a) lower alkyl which may be substituted with 5 one or more substituent(s) selected from the group consisting of cycloalkyl, aryl which may be further substituted with 10 aryl(s), and heterocyclyl, (b) lower alkenyl which may be substituted with one or more substituent(s) selected from the group consisting of aryl and heterocyclyl, 15 (c) cycloalkyl, (d) aryl which may be substituted with one or more substituent(s) selected from the group consisting of lower alkyl, 20 aryl which may be further substituted with hydroxy(s), lower alkoxy, aryloxy, hydroxy, and 25 halogen, (e) heterocyclyl which may be substituted with one or more substituent(s) selected from the group consisting of lower alkyl, 30 aryl which may be further substituted with halogen(s), and halogen, (f) aryloxy, or (g) amino which may be substituted with aryl(s) 35 which may be substituted with one or more

substituent(s) selected from the group consisting of aryl and heterocyclyll;

or a pharmaceutically acceptable salt thereof,

5

comprising, reacting a compound (IIa):

O OH
$$Y = N R^{6}$$

$$H$$
(II a)

(wherein Y and R^6 are each as defined above), or its reactive derivative at the carboxy group or the salt thereof, with a resin-bound compound (IIIc):

$$R^{2'}$$
 O \mathbb{P} O \mathbb{P} O \mathbb{P} O \mathbb{P}

(wherein R^{2} is as defined above, and Θ is polymer), or its reactive derivative at the amino group or the salt thereof to give a compound (IVc):

15

(wherein Y, $\ensuremath{\mathbb{P}}$ $\ensuremath{\mathbb{R}}^2$ and $\ensuremath{\mathbb{R}}^6$ are as defined above), or its

salt; .

reacting the compound (IVc):

$$\begin{array}{c|c}
O & H \\
N & O \\
P & O
\end{array}$$

$$\begin{array}{c|c}
N & R^6 \\
H & (IV_C)
\end{array}$$

5 (wherein Y, \bigcirc R² and R⁶ are as defined above), or its salt, with a compound (V):

$$R^7$$
 OH (V)

(wherein Q and R^7 are as defined above), or its reactive derivative at the carboxy group (in case of Q is -CO-)/the sulfo group (in case of Q is $-SO_2-$), or the salt thereof to give a compound (Ia-2'):

O
$$\mathbb{R}^{2^{2}}$$
 O $\mathbb{P}^{2^{2}}$ O $\mathbb{P}^{2^{2}}$ (I a-2')

(wherein Q, Y, \mathbb{P} , \mathbb{R}^2 ', \mathbb{R}^6 , and \mathbb{R}^7 are as defined above), or its salt; and

subjecting the compound (Ia-2'):

15

$$\begin{array}{c|c}
 & H \\
 & N \\
 & R^{2'} & O
\end{array}$$

$$\begin{array}{c}
 & NR^6 \\
 & Q \\
 & R^7 \\
 & (I a-2')
\end{array}$$

(wherein Q, Y, \mathbf{P} , \mathbf{R}^2 ', \mathbf{R}^6 , and \mathbf{R}^7 are as defined above), or its salt to a cleavage reaction of the resin.

5 9. A compound of any one of Claims 1 to 5 for use as a medicament.

10

- 10. The compound of Claim 9 for use in the treatment and/or prevention of PGE_2 mediated diseases in human beings or animals.
- 11. A medicament comprising a compound of any one of Claims 1 to 5 as an active ingredient.
- 15 12. A pharmaceutical composition comprising a compound of any one of Claims 1 to 5 as an active ingredient, in association with a pharmaceutically acceptable carrier or excipient.
- $20\,$ 13. An agonist or antagonist of PGE $_2$ consisting of a compound of any one of Claims 1 to 5.
- 14. A method for treatment and/or prevention of PGE $_2$ mediated diseases which comprises administering an effective amount of the compound of any one of Claims 1 to 5 to human beings or animals.

15. A method for treating or preventing kidney dysfunction, inflammatory conditions, various pains, collagen diseases, autoimmune diseases, various immunity diseases, analgesic, thrombosis, allergic disease, cancer or neurodegenerative diseases which comprises administering an effective amount of a compound of any one of Claims 1 to 5 to human beings or animals.

5

15

- 10 16. Use of a compound of any one of Claims 1 to 5 as a medicament.
 - 17. Use of a compound of any one of Claims 1 to 5 as an agonist or an antagonist of PGE_2 -sensitive receptor.
- 18. Use of the compound of any one of Claims 1 to 5 for treatment and/or prevention of PGE_2 mediated diseases in human beings or animals.
- 20 19. A commercial package comprising the pharmaceutical composition containing the compound identified in any one of any one of Claims 1 to 5 and a written matter associated therewith, wherein the written matter states that the compound (I) can or should be used for preventing or treating PGE2 mediated diseases.